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I hereby certify under 37 C.F.R. § 1.8(a) that this correspondence is being deposited with the United States Postal Service as first class mail with sufficient postage on the date indicated above and is addressed to Mail Stop AF, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313/1450.

Tracey Simmons

Printed name of person mailing correspondence

Signature of person mailing correspondence

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant:

Roy A. Gravel et al.

Art Unit:

1632

Serial No.:

09/487,841

Examiner:

Shin-Lin Chen

Filed:

January 19, 2000

Customer No.:

21559

Title:

HUMAN METHIONINE SYNTHASE REDUCTASE: CLONING,

AND METHODS FOR EVALUATING RISK OF, PREVENTING, OR TREATING NEURAL TUBE DEFECTS, CARDIOVASCULAR

DISEASE, CANCER, AND DOWN'S SYNDROME

Mail Stop AF Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

STATEMENT UNDER 37 C.F.R. § 1.821

Enclosed is a Sequence Listing in accordance with the requirements of 37 C.F.R. §§ 1.821 through 1.825 and consisting of 28 pages.

As required by 37 C.F.R. § 1.821(c), the Sequence Listing appears as a separate part of the application. Each sequence in the application appears separately in the Sequence Listing, and each sequence in the Sequence Listing is assigned a separate sequence identifier.

As required by 37 C.F.R. § 1.821(d), the sequence identifiers are used throughout

the application description and claims to refer to their respective sequences.

As required by 37 C.F.R. § 1.821(e), enclosed is a diskette containing a copy of the Sequence Listing in computer readable form.

As required by 37 C.F.R. § 1.821(f), I hereby state that the contents of the computer readable form of the Sequence Listing are the same as the contents of the paper copy.

As required by 37 C.F.R. § 1.821(g), I hereby state that this submission contains no new matter.

If there are any charges or any credits, please apply them to Deposit Account No. 03-2095.

Respectfully submitted,

Date: Scenber 15, Woy

Kristina Bieker-Brady, Ph.D., P.C.

Reg. No. 39,109

Clark & Elbing LLP 101 Federal Street Boston, MA 02110

Telephone: 617-428-0200 Facsimile: 617-428-7045



SEQUENCE LISTING

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Leclerc, Daniel
Wilson, Aaron
Rosenblatt, David

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Glu	Glu 50	Val	Pro	Glu	Phe	Thr 55	Lys	Ile	Gln	Thr	Leu 60	Thr	Ser	Ser	Val
Arg 65	Glu	Ser	Ser	Phe	Val 70	Glu	Lys	Met	Lys	Lys 75	Thr	Gly	Arg	Asn	Ile 80
Ile	Val	Phe	Tyr	Gly 85	Ser	Gln	Thr	Gly	Thr 90	Ala	Glu	Glu	Phe	Ala 95	Asn
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90

75

55

70

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Gly Leu Trp Pro Ala Leu Arg Lys His Phe Arg Ser Ser Arg Gly Gln
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Glu Glu Ile Ser Gly Ala Leu Pro Val Ala Ser Pro Ala Ser Leu Arq
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Thr Asp Leu Val Lys Ser Glu Leu Leu His Ile Glu Ser Gln Val Glu
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Gln Asn Ala Val Asn Ser Asn Gln Ser Asn Val Val Ile Glu Asp Phe
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Gln Pro Arg Pro Tyr Ser Cys Ala Ser Ser Ser Leu Phe His Pro Gly
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Lys Leu His Phe Val Phe Asn Ile Val Glu Phe Leu Ser Thr Ala Thr
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Val Ala Ser Val Leu Gln Pro Asn Ile His Ala Ser His Glu Asp Ser
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Leu Gln Glu Leu Cys Ser Lys Gln Gly Ala Ala Asp Tyr Ser Arg Phe
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Gln Pro Arg Pro Tyr Ser Cys Ala Ser Ser Ser Leu Phe His Pro Gly
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Lys Leu His Phe Val Phe Asn Ile Val Glu Phe Leu Ser Thr Ala Thr
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Thr Glu Val Leu Arg Lys Gly Val Cys Thr Gly Trp Leu Ala Leu Leu
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Val Ala Ser Val Leu Gln Pro Asn Ile His Ala Ser His Glu Asp Ser
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Pro Gly Thr Gly Ile Ala Pro Phe Ile Gly Phe Leu Gln His Arg Glu
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Lys Leu Gln Glu Gln His Pro Asp Gly Asn Phe Gly Ala Met Trp Leu
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Phe Phe Gly Cys Arg His Lys Asp Arg Asp Tyr Leu Phe Arg Lys Glu
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Leu Arg His Phe Leu Lys His Gly Ile Leu Thr His Leu Lys Val Ser
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Val Gln Asp Asn Ile Gln Leu His Gly Gln Gln Val Ala Arg Ile Leu
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Leu Gln Glu Asn Gly His Ile Tyr Val Cys Gly Asp Ala Lys Asn Met
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<213> Homo sapiens
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<210> 46 <211> 697 <212> PRT

<213> Homo sapiens

<400> 46

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Gln	Asn 210	Ala	Val	Asn	Ser	Asn 215	Gln	Ser	Asn	Val	Val 220	Ile	Glu	Asp	Phe
Glu 225	Ser	Ser	Leu	Thr	Arg 230	Ser	Val	Pro	Pro	Leu 235	Ser	Gln	Ala	Ser	Leu 240
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			260					265				Ser	270	-	
		275					280					Thr 285			_
	290					295				_	300	Ser			_
305					310	_				315		Cys			320
				325					330			Leu		335	_
			340					345				Thr	350		
		355					360			_	_	Ser 365 Lys			
	370					375					380	Буs			
385					390					395		Tyr			400
				405					410		_	Leu		415	
			420					425	_			Leu	430		
		435					440					445 Phe		-	
	450					455					460	Ser			-
465					470					475		Leu			480
				485					490			His		495	
			500					505				Arg	510	-	
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Pro	530 Gly	Thr	Gly	Ile	Ala	535 Pro	Phe	Ile	Gly	Phe	540 Leu	Gln	His	Arg	Glu
545 Lys	Leu	Gln	Glu	Gln	550 His	Pro	Asp	Gly	Asn	555 Phe	Gly	Ala	Met	Trp	560 Phe
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Arg	His		580 Leu	Lys	His	Gly		585 Leu	Thr	His	Leu	Lys	590 Val	Ser	Phe
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<210> 47 <211> 2093 <212> DNA <213> Homo sapiens

<400> 47

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<210> 48 <211> 689

<212> PRT

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Leu	His	Phe	Val	Phe	Asn	Ile	Val	Glu	Phe	Leu	Ser	Thr	Ala	Thr	Thr	
465	** 7	_	_	_	470	** 7	~	 1	~ 3	475	.		- .	.	480	
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Tlo	Cox	Tou	580	Mot	G1.,	Cor	T 011	585	7 ~~	Dho	Dwo	Con	590 Gln	~ 1	Mot	
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	Ile	Phe	Met	Cys 645		Glu	Met	Gln	Arg 650		Trp	Pro	Arg	Met 655		
Met	Met	Pro			Lys	Ala	Lys	_		Glu	Leu	Lys	Asn		Gln	
Lvs	Pro	Tro	660 Pro	ī.en	Lvs	Lvs	Lvs	665 Asn	Δla	Thr	Dhe	Δra	670 Ile	Dhe	Glv	
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Lys Arg Tyr Leu Gln Asp Ile Trp Ser
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